



Year: 2000

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Abstract: Objectives: Magnetic resonance (MRI) velocity mapping was used to evaluate non-invasively the flow profiles of the ascending aorta in normal volunteers and in patients with an aortic (mechanical) valve prosthesis. Background: In patients with artificial aortic valves the flow profile in the ascending aorta is severely altered. These changes have been associated with an increased risk of thrombus formation and mechanical hemolysis. Methods: Velocity profiles were determined 30 mm distal to the aortic valve in six healthy volunteers and seven patients with aortic valve replacement (replacement within the last 2 years) using ECG triggered phase contrast MRI. Peak flow, mean flow and mean reverse flow were measured in intervals of 25 ms during the entire heart cycle. Systolic reverse flow, end-systolic closing and diastolic leakage volume were calculated for all subjects. Results: Peak flow velocity during mid-systole was significantly higher in patients with valvular prosthesis than in normals (mean \pm SD, 1.9 ± 0.4 m/s vs. 1.2 ± 0.03 m/s, $P<0.001$) with a double peak and a zone of reversed flow close to the inner (left lateral) wall of the ascending aorta of the patients. Closing volume was significantly larger in patients than in controls (-3.3 ± 1.2 ml/beat vs. -0.9 ± 0.5 ml/beat; $P<0.001$). There was reverse flow during systole in valvular patients amounting to $15.7\pm 6.7\%$ of total cardiac output compared to $2.3\pm 1.2\%$ in controls ($P<0.001$). Diastolic mean flow was negative in patients after valve replacement but not in controls (-11.0 ± 15.2 ml/beat vs. 6.8 ± 3.2 ml/beat; $P<0.01$). Conclusions: The following three major quantitative observations have been made in the present study: (1) Mechanical valve prostheses have an increased peak flow velocity with a systolic reverse flow at the inner (left lateral) wall of the ascending aorta. (2) A double peak flow velocity pattern can be observed in patients with bileaflet (mechanical) prosthesis. (3) The blood volume required for leaflet closure and the diastolic leakage blood volume are significantly higher for the examined bileaflet valve than for native heart valves

DOI: <https://doi.org/10.1007/bf02613108>

Posted at the Zurich Open Repository and Archive, University of Zurich

ZORA URL: <https://doi.org/10.5167/uzh-156314>

Journal Article

Published Version

Originally published at:

Botnar, René; Nagel, Eike; Scheidegger, Markus Beat; Pedersen, Erik Morre; Hess, Otto; Boesiger, Peter (2000). Assessment of prosthetic aortic valve performance by magnetic resonance velocity imaging. *Magma*, 10(1):18-26.

DOI: <https://doi.org/10.1007/bf02613108>

Assessment of prosthetic aortic valve performance by magnetic resonance velocity imaging

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Received 17 May 1999; received in revised form 13 August 1999; accepted 26 August 1999

Abstract

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Results: Peak flow velocity during mid-systole was significantly higher in patients with valvular prosthesis than in normals (mean \pm SD, 1.9 ± 0.4 m/s vs. 1.2 ± 0.03 m/s, $P < 0.001$) with a double peak and a zone of reversed flow close to the inner (left lateral) wall of the ascending aorta of the patients. Closing volume was significantly larger in patients than in controls (-3.3 ± 1.2 ml/beat vs. -0.9 ± 0.5 ml/beat; $P < 0.001$). There was reverse flow during systole in valvular patients amounting to $15.7 \pm 6.7\%$ of total cardiac output compared to $2.3 \pm 1.2\%$ in controls ($P < 0.001$). Diastolic mean flow was negative in patients after valve replacement but not in controls (-11.0 ± 15.2 ml/beat vs. 6.8 ± 3.2 ml/beat; $P < 0.01$).

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Keywords: Aortic flow; Magnetic resonance velocity mapping; Prosthetic cardiac valves; Bileaflet valves; Reverse flow; Aortic stenosis

1. Introduction

Since the first use of cardiac valve prostheses in 1951 by Charles Hufnagel more than 50 different valve prostheses have been implanted. Today cardiac valve replace-

ment has become a routine procedure (worldwide 175 000 per year), but there are still unsolved long-term issues. The most severe problems are thrombosis, hemolysis and tissue overgrowth [1–4]. These problems are mainly dependent on valve design, blood flow velocity and turbulence fields [5–11]. Regions of slow flow, reversed flow and flow separation may promote formation of blood clots, leading to thrombus formation and cerebrovascular embolization [12]. Anticoagulation greatly reduces the incidence of thromboembolism, but is associated with an increased risk of hemorrhage.

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The purpose of the present study was (1) to investigate the utility of magnetic resonance (MR) flow velocity mapping for measuring blood flow patterns in the ascending aorta of patients with an aortic valve prosthesis; and (2) to study the differences in velocity and flow patterns between these patients and a group of volunteers with healthy native valves. The presence of abnormal flow patterns distal to different types of mechanical valve prostheses has been described both under in-vitro conditions and in experimental animals [10,13–19]. Due to the complexity of the flow velocity fields distal to a prosthetic valve and lack of accurate non-invasive measurement techniques, precise measurements are difficult to obtain and only a few studies have been performed in humans [20–23].

In the present study a short echo-time MRI-technique was applied [24–27]. It allows for the assessment of blood flow velocities across the entire area of the ascending aorta with high temporal and spatial resolution. We hypothesize that the flow patterns among those with native valves represent aortic flow under normal physiologic conditions. The performance of a valvular prosthesis should therefore be optimal if similar flow patterns are observed.

2. Materials and methods

2.1. Materials

We studied six healthy adult volunteers (male, mean age, 27 years, range, 25–28) and seven patients (male, mean age, 66 years, range, 61–71) with bileaflet aortic prosthesis implanted 1–2 years prior to study. All patients were in stable condition with no evidence of valve dysfunction on echocardiographic examinations. No patient had symptoms suggestive of coronary artery disease and all had no evidence of disease on coronary angiograms prior to surgery. In the patient collective mean systolic blood pressure was $138 \text{ mmHg} \pm 16 \text{ mmHg}$ and mean heart rate was $71 \pm 10 \text{ bpm}$ (mean \pm SD).

2.2. Magnetic resonance velocity mapping

Written informed consent was obtained from all subjects. Measurements were performed on a 1.5 Tesla Philips NT whole body scanner. The subjects were examined in prone position with a prototype cardiac receiver coil. An ECG gated Turbo Field Echo (TFE)

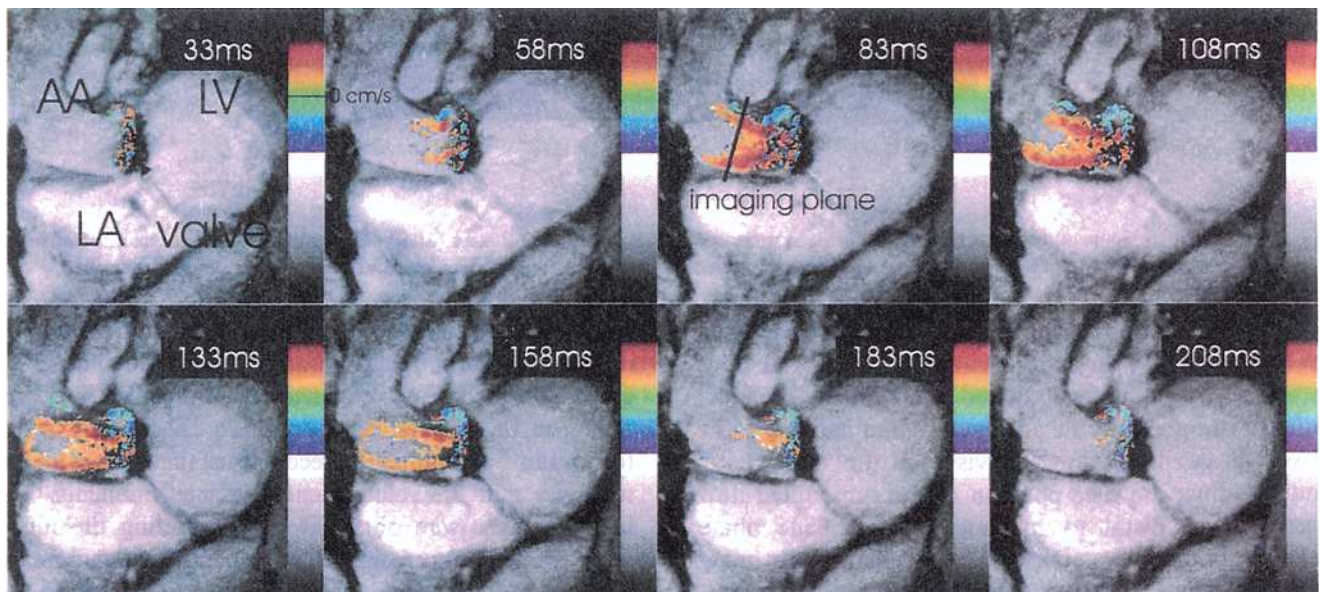


Fig. 1. Color-coded velocity images of the aortic outflow tract during systolic contraction in a patient after aortic valve replacement. The mechanical bileaflet valve causes a disk shaped black artifact. An imaging plane through the left ventricle (LV), left atrium (LA) and ascending aorta (AA) has been chosen for flow visualization. Temporal resolution is 50 ms per frame. The image plane of the subsequent through plane flow measurement is indicated as white bar. Red colors stand for antegrade, blue for retrograde flow.

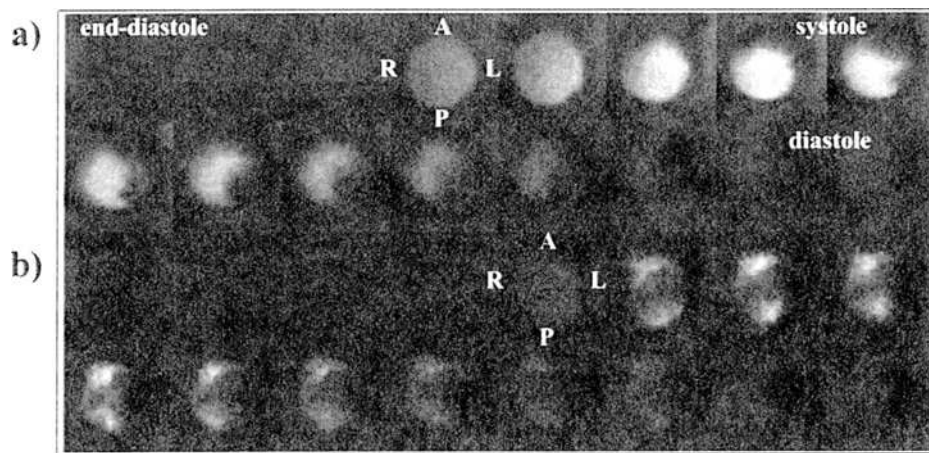


Fig. 2. Axial flow velocity images through the ascending aorta 30 mm above the aortic annulus. A normal volunteer (a) and a patient with mechanical valve prosthesis of the bileaflet type (b) are shown during systolic ejection. The brightness of the images is proportional to forward flow velocities the darkness to retrograde flow velocities. Temporal resolution is 25 ms per frame. R: right, L: left, A: anterior, P: posterior.

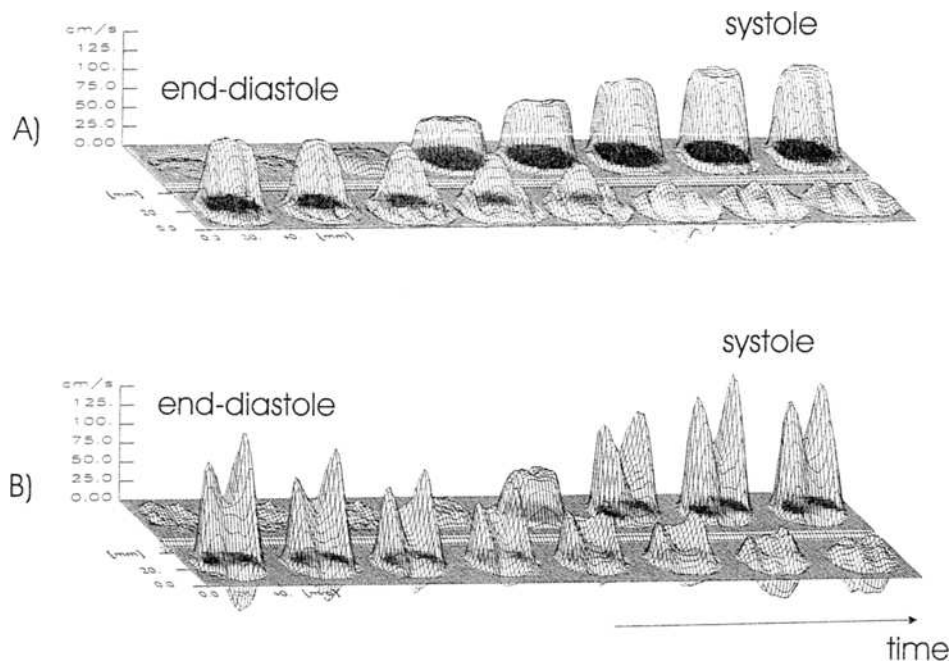


Fig. 3. Flow velocity maps in the ascending aorta during systolic ejection in a normal volunteer (a) and in a patient with a mechanical valve (b). a, b) Top left frame: end-diastole; top right frame: systole, bottom right frame: early diastole. Temporal resolution is 25 ms per frame.

sequence with three orthogonal slice packages was used to determine the orientation of the aortic valve or prosthesis. A double-angulated TFE image was acquired orthogonal to the aortic valve and parallel to the proximal ascending aorta to visualize the velocity jets and to adjust the slice position of the subsequent flow measurement distal to the aortic valve. Ciné phase velocity mapping was performed parallel to this imaging plane (Fig. 1) and orthogonal (Fig. 2) to the ascending aorta approximately 30 mm (1 annulus diameter) distal to the aortic valve.

To reduce image artifacts due to local blood flow acceleration and turbulence, a short echo time (TE =

2.8ms (13 mT/m, 45 mT/m/ms)–6.4 ms (10 mT/m, 10 mT/m/ms)) partial echo (62.5% of the full echo) ECG gated phase contrast sequence was applied. After zero filling of the missing echo samples a 2D-Fourier transformation was applied to reconstruct the images of the reference and the velocity encoded measurements. Velocity images were obtained by subtracting the reference from the corresponding velocity encoded phase images. Velocity encoding was set according to the maximal expected blood flow velocity and varied between 120–280 cm/s. In-plane velocity maps were used to visualize the velocity jets originating from the different orifices of the prosthetic valves along the course of

the aorta (Fig. 1). Through-plane velocity images allowed distinguishing between antegrade and retrograde flow (Figs. 2 and 3).

The slice thickness was 10 mm, field of view $350 \times 280 \text{ mm}^2$ and image matrix 256×204 resulting in an in-plane resolution of $1.37 \times 1.37 \text{ mm}^2$. The time resolution was 25 ms and the echo time varied between 3 and 6 ms. Velocity compensated and velocity encoded measurements were performed in consecutive heart phase intervals. Acquisition time was 5.8 min for a heart rate of 70 beats/min.

2.3. Velocity data analysis

Data analysis was performed on a DEC-Alpha (Digital Equipment Corporation, USA) workstation with a flow analysis software package developed at our institution. Vessel contours are either drawn manually or

using movable and deformable ellipses. The contour identification can either be done on the modulus or the velocity (phase) images of each heart phase. The obtained contours are copied to the corresponding modulus or velocity (phase) images. After vessel wall segmentation through-plane mean flow (Fig. 4), peak velocity (Fig. 5), mean forward and mean reverse flow (Fig. 6) are calculated for each heart phase. Peak flow velocity was defined for each heart phase as the highest velocity value measured within the lumen of the aorta. Mean flow rate per second is defined as sum of mean forward and mean reverse flow rate. Linear interpolation was used to estimate mean flow rates per beat. To allow proper comparison of different subjects the time axis is normalized by defining end-systole (100%) as the time of the zero crossing of mean flow rate (Fig. 4). Using the approach of Yoganathan, closing and leakage volume were defined [28]. Integration of the mean

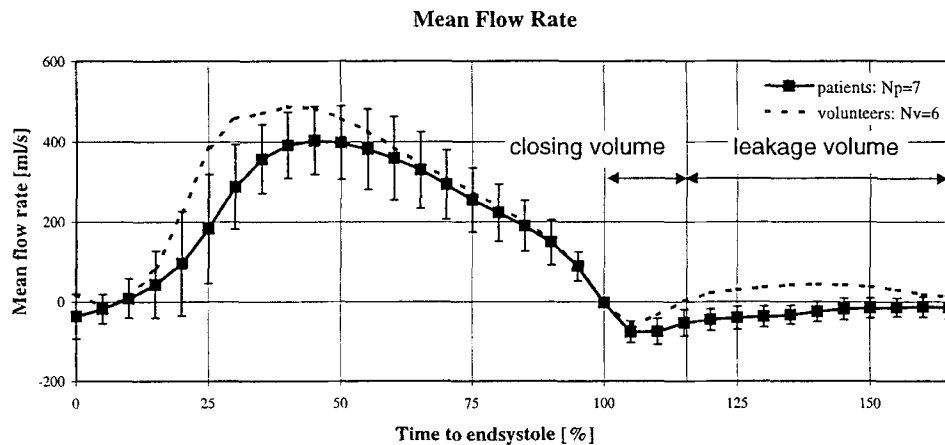


Fig. 4. Mean flow rate (\pm standard deviation) over time in six volunteers (dashed) and seven patients after aortic valve replacement (line). During systole no significant differences in mean flow rate can be observed but during diastole flow is significantly lower in patients and becomes even negative due to leakage flow. Definitions of closing and leakage volume are given accordingly to Yoganathan [28]. X-axis: time normalized for end-systole; Y-axis: mean flow rate.

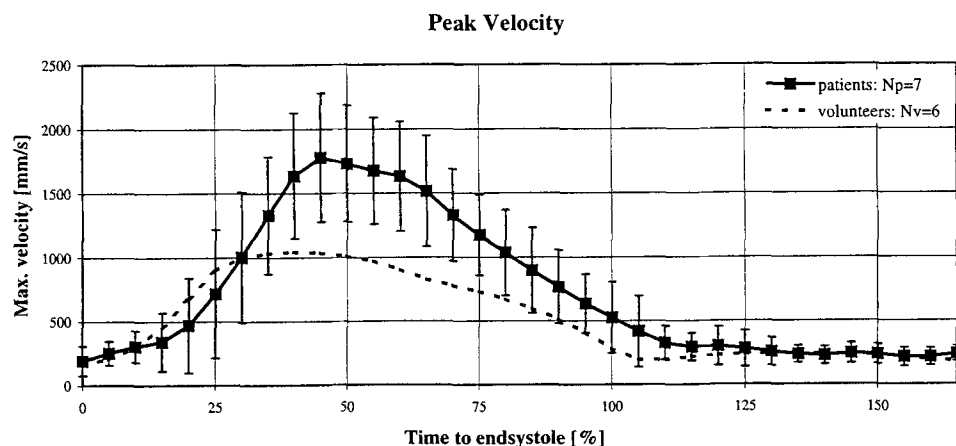


Fig. 5. Peak flow velocity (\pm standard deviation) in six volunteers (dashed) and seven patients with a mechanical bileaflet valve (line). During early systole peak flow is significantly higher in patients with valvular prosthesis than in controls probably due to the smaller valve orifice. Diastolic peak flow velocities are similar in the two groups. X-axis: time normalized for end-systole; Y-axis: peak flow velocity.

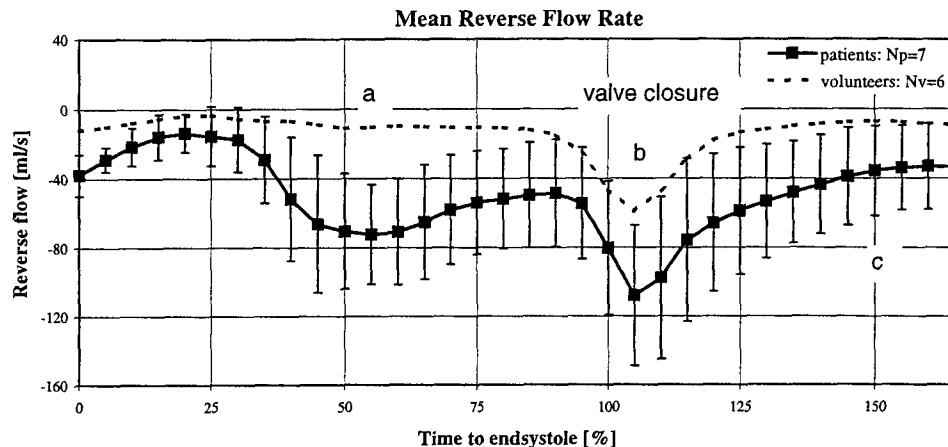


Fig. 6. Mean reverse flow-rate (\pm standard deviation) over time in six volunteers (dashed) and seven patients with mechanical bileaflet valves (line). In controls only a small reverse flow can be seen during systolic ejection with a large peak during valve closure (end-systole; 100%). In contrast, valvular patients show a large reverse flow during systolic ejection with a peak during valve closure. X-axis: time normalized for end-systole; Y-axis: mean reverse flow rate.

flow rate from 100 to 115% of end-systole was defined as closing volume and the integral from 115% of end-systole to the next R-wave as diastolic flow volume. The velocity maps are visualized as 3D-mesh plots with the underlying corresponding gray-value coded velocity map on a high resolution video monitor (Fig. 3).

2.4. Statistics

All data are represented as mean \pm SD. The unpaired Student's *t*-test was applied for comparisons of mean values between patients and normal volunteers. Significance was defined by a *P* value of ≤ 0.05 .

3. Results

All patients and volunteers completed MR studies without incidence. Representative in-plane and through-plane velocity images of a volunteer and a patient are shown in Fig. 1 and Fig. 2, respectively. The quality of the phase maps (Fig. 2) suggest that no significant flow voids were present. A horizontal imaging plane through the left ventricle (LV), left atrium (LA) and ascending aorta (AA) has been chosen for flow visualization (Fig. 1). The valve prosthesis can be observed as a black disk. The flow velocities distal to the prosthetic valve are displayed as color encoded images overlaid on an anatomical background image of the aortic outflow tract. Three antegrade velocity jets, two marginal and one central are visible throughout systole. During early systole, the marginal jets follow the curvature of the aorta. Then, from mid-systole to end-systole, the two outer jets run parallel to each other and are relatively orthogonal to the valvular plane. Due to susceptibility artifacts of the prosthesis ring, flow

velocities are noisy in the vicinity of the valve.

A 3D-reconstruction of the flow velocities in the ascending aorta is depicted in Fig. 3 for a normal volunteer and a patient with bileaflet prosthesis. In the normal subject, a flat flow profile can be observed during systolic ejection with some flow reversed during early diastole. In contrast, the patient with mechanical valve shows two large peaks during systolic ejection with simultaneous systolic backward flow and large early diastolic reverse flow (= closing volume).

3.1. Flow velocity data

Mean flow rates are calculated by integration of flow velocities over the vessel areas (Fig. 4). From end-diastole to end-systole no significant difference in mean flow rate can be observed in patients with mechanical valves. A significant difference in mean flow rate can be observed during valve closure (-3.3 ± 1.2 ml/beat vs. -0.9 ± 0.5 ml/beat; $P < 0.001$) and during diastole (-11.0 ± 15.2 ml/beat vs. $+6.8 \pm 3.2$ ml/beat; $P < 0.01$) revealing slight leakage flow in patients with a valvular prosthesis. The measured values for each subject are presented in Table 1.

Temporal changes in peak flow velocity are similar during early systole (Fig. 5). However, during systolic ejection significantly higher peak flow velocities (1.9 ± 0.4 m/s vs. 1.2 ± 0.03 m/s; $P < 0.001$) can be observed in patients with mechanical valves than in controls. During early and mid-diastole, peak flow velocities are similar for valvular patients and controls ($P > 0.2$).

3.2. Retrograde flow

There were significant differences in retrograde flow between patients and controls (Fig. 6) during mid-sys

tole (a) (-72 ± 29 ml/s vs. -11 ± 11 ml/s; $P < 0.001$) and also during diastole (c) (-36 ± 26 ml/s vs. -7 ± 9 ml/s; $P < 0.04$). A tendency of increased reverse flow volume was observed during end-systole (b) (-108 ± 41 ml/s vs. -59 ± 26 ml/s; $P < 0.07$) (valve closure).

3.3. Aortic diameter

The diameter of the aorta was significantly higher in valvular patients (36 ± 2 mm vs. 30 ± 2 mm, $P < 0.001$).

4. Discussion and conclusions

An increasing number of patients require aortic valve replacement because of degenerative aortic valve disease. Non-invasive assessment and follow up of valve performance is, therefore, important to detect valve dysfunction in appropriate time. Doppler echocardiography represents currently the gold standard for detection of valve dysfunction but newer techniques such as MR imaging may provide more detailed information with regard to flow distribution, presence or absence of antegrade or retrograde flow jets as well as anatomy of the ascending aorta.

Previous hemodynamic examinations of artificial heart valves have used echocardiographic methods for diagnosis of valvular dysfunction [28–30]. However, these studies have so far been limited to one-dimensional flow profiles, except from perioperative studies [31], and the results may be observer dependent with variable image

quality and interobserver variability. For the description of the mechanical function of valve prostheses quantitative data on ante- and retrograde blood flow volumes may provide additional information.

Compared to echocardiography, MRI blood flow measurements not only allow determination of regional blood flow velocities, but also overall blood flow rate through the aortic valve. Therefore, accurate predictions can be obtained on ante- and retrograde flow, closing and leakage blood volumes as well as systolic and diastolic mean and peak flow velocities.

This study demonstrates that using a short echo time phase velocity mapping technique, it is possible to obtain quantitative information about the blood velocity distribution across the whole vessel area downstream of artificial mechanical heart valves. The measurements can be obtained with a high spatial (1.37×1.37 mm²) and temporal (25 ms) resolution. Only neglectable flow voids due to turbulence known to be present downstream of artificial heart valves also in-vivo [16,20] were seen. This is in accordance with recent in-vitro studies of velocity fields at artificial heart valves using magnetic resonance phase velocity mapping [18,19].

4.1. Limitations

In this study only patients without signs of valvular dysfunction were examined. The usefulness of the presented technique for the assessment of valve dysfunction therefore still has to be proven. However, it can be expected that for mild and moderate aortic stenoses, this

Table 1

Values are mean \pm SD of the volunteer group and the patient group for closing volume, diastolic flow volume (leakage volume), systolic reverse flow volume and aortic diameter^a

Patient	Closing volume (ml/beat)	Diastolic flow volume (leakage volume) (ml beat)	Mean aortic diameter (mm)
1	-1.6	-1.7	35
2	-2.6	-4.9	38
3	-3.5	-37.0	32
4	-2.2	0.0	37
5	-3.8	-2.1	38
6	-4.7	-3.4	33
7	-4.6	-28.6	37
Mean	-3.3	-11.0	36.0
SD	1.2	15.2	2
Volunteer			
1	-0.8	12.2	32
2	-0.03	5.6	28
3	-1.3	5.8	27
4	-1.4	5.2	32
5	-1.3	8.7	31
6	-0.6	3.3	31
Mean	-0.9	6.8	30
SD	0.5	3.2	2
P value	<0.001	<0.01	<0.001

^a All quantities were quantified 30 mm distal to the aortic valve, and the prosthetic aortic valve, respectively. Instead of leakage volume, diastolic flow volume was quantified in volunteers. Leakage was only observed in patients with prosthetic valves.

technique still provides reliable phase maps and therefore information about the hemodynamic performance of valvular prosthesis.

All examined patients had surgery 1–2 years prior to the MR examination. This short time span therefore did not allow studying aging effects of prosthetic valves. By selecting another patient population these effects, however, should be accessible with the presented technique.

The patient and the volunteer population of the presented study were not age matched. This might cause a bias in the comparison of the both groups. The finding that MR allows for the assessment of flow fields (jet like flow, closing volume, leakage volume) distal to prosthetic valves, however, should not be influenced by this fact.

MRI-velocity mapping is furthermore limited to a certain extent by image artifacts, which are caused by turbulent blood flow, and in-plane flow [32–34]. Reliable images can be acquired with very short echo times as they can be achieved with new MR-scanners using partial echo acquisition scheme like the one presented here. The metallic ring of the artificial valve induces severe image artifacts and, thus, flow velocities can be assessed only in a certain distance from the valve. In-vitro studies carried out in our lab showed that reliable flow measurements still can be performed in a distance of a $\frac{1}{4}$ of the diameter of the valve [35]. At this location even the central high velocity jet, which is assessed by Doppler echocardiography to estimate the pressure drop, could be assessed. In-vivo measurements, however, are subject to valvular motion, which, if not compensated for can cause artifacts and phase drifts due to the metallic ring of the prosthesis. Kozerke and co-workers [36] therefore suggested adapting the slice position for each single heart phase. In the present study this technique was not incorporated and therefore it only allows for aortic flow measurements at a certain distance to the aortic valve plane ($> 15\text{mm}$). Because of this limitation the central peak velocity jet that is assessed by US cannot be measured and one is limited to the assessment of the two outer velocity jets that remain visible also further downstream. Kozerke and co-workers [36] also showed that without valvular motion correction early diastolic retrograde flow is overestimated by about 20%. This number might be, however, smaller in patients with aortic regurgitation where overall diastolic retrograde flow is increased.

Since the slice location is distal to the coronary ostia, coronary blood flow and aortic compliance might cause a bias of the results as described by Chatzimavroudis and co-workers [37]. Incorporating slice tracking in future measurements, however, should compensate for these errors. Prospective triggering does not allow the assessment of late diastole unless the measurement is performed over two heartbeats. This seems, however,

only to be a small limitation since diastolic blood flow is slow with only small temporal variations. Extrapolation of diastolic flow should therefore only result in minor errors. Temporal resolution of the MR measurements is less than for Doppler echocardiography but likely adequate for assessing valvular function in the clinical setting.

4.2. Clinical aspects

High velocity jets, as they are measured in artificial heart valves, but do not occur in healthy natural valves, are associated with turbulent blood flow. These velocity jets may increase regional wall shear stress and favour dilatation as well as local aneurysm formation. This may be more severe if the valve is not properly aligned and the jet is directed towards the wall. Proper alignment of the valve prosthesis within the aorta and appropriate sizing of the valve are, therefore, of clinical importance to reduce regional wall stress and aneurysm formation.

Clinical evaluation of artificial heart valves by MRI might thus become more important since these velocity maps can be achieved non-invasively. From these measurements, information on valve function, prosthesis alignment and presence of transvalvular or paravalvular jets can be obtained. This high degree of detail might allow the disclosure of even minor quantitative hemodynamic changes over time in individual patients. Further repeatability studies are needed to clarify these issues.

The following three major quantitative observations have been made in the present study: (1) Mechanical aortic valve prostheses induce an increased peak flow velocity with a systolic reverse flow at the inner (left lateral) wall of the ascending aorta. (2) A double peak flow velocity pattern is observed in patients with bileaflet (mechanical) prosthesis. (3) The blood volume for leaflet closure and the diastolic leakage volume are significantly higher than blood volume for leaflet closure in native aortic valves.

In conclusion magnetic resonance flow velocity mapping is a promising tool for a detailed, quantitative non-invasive assessment of the hemodynamic valvular performance in patients with valve prostheses.

Acknowledgements

We acknowledge the support of EUREKA and the Swiss Commission for Technology and Innovation; Philips Medical Systems, Best, The Netherlands; Desirée and Niels Foundation, Denmark; The Danish Heart Foundation and Sulzer Innotec, Winterthur, Switzerland. We also like to thank Dr Warren Manning for helpful discussions.

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